Application No: 10/527,716

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| | | |
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| NEWS 1 | | Web Page for STN Seminar Schedule - N. America |
| NEWS 2 | JAN 08 | CHEMLIST enhanced with New Zealand Inventory of Chemicals |
| NEWS 3 | JAN 16 | CA/CAplus Company Name Thesaurus enhanced and reloaded |
| NEWS 4 | JAN 16 | IPC version 2007.01 thesaurus available on STN |
| NEWS 5 | JAN 16 | WPIDS/WPINDEX/WPIX enhanced with IPC 8 reclassification data |
| NEWS 6 | JAN 22 | CA/CAplus updated with revised CAS roles |
| NEWS 7 | JAN 22 | CA/CAplus enhanced with patent applications from India |
| NEWS 8 | JAN 29 | PHAR reloaded with new search and display fields |
| NEWS 9 | JAN 29 | CAS Registry Number crossover limit increased to 300,000 in multiple databases |
| NEWS 10 | FEB 15 | PATDPASPC enhanced with Drug Approval numbers |
| NEWS 11 | FEB 15 | RUSSIAPAT enhanced with pre-1994 records |
| NEWS 12 | FEB 23 | KOREAPAT enhanced with IPC 8 features and functionality |
| NEWS 13 | FEB 26 | MEDLINE reloaded with enhancements |
| NEWS 14 | FEB 26 | EMBASE enhanced with Clinical Trial Number field |
| NEWS 15 | FEB 26 | TOXCENTER enhanced with reloaded MEDLINE |
| NEWS 16 | FEB 26 | IFICDB/IFIPAT/IFIUDB reloaded with enhancements |
| NEWS 17 | FEB 26 | CAS Registry Number crossover limit increased from 10,000 to 300,000 in multiple databases |
| NEWS 18 | MAR 15 | WPIDS/WPIX enhanced with new FRAGHITSTR display format |
| NEWS 19 | MAR 16 | CASREACT coverage extended |
| NEWS 20 | MAR 20 | MARPAT now updated daily |
| NEWS 21 | MAR 22 | LWPI reloaded |
| NEWS 22 | MAR 30 | RDISCLOSURE reloaded with enhancements |
| NEWS 23 | APR 02 | JICST-EPLUS removed from database clusters and STN |
| NEWS 24 | APR 30 | GENBANK reloaded and enhanced with Genome Project ID field |
| NEWS 25 | APR 30 | CHEMCATS enhanced with 1.2 million new records |
| NEWS 26 | APR 30 | CA/CAplus enhanced with 1870-1889 U.S. patent records |
| NEWS 27 | APR 30 | INPADOC replaced by INPADOCDB on STN |
| NEWS 28 | MAY 01 | New CAS web site launched |
| NEWS 29 | MAY 08 | CA/CAplus Indian patent publication number format defined |
| NEWS 30 | MAY 14 | RDISCLOSURE on STN Easy enhanced with new search and display fields |
| NEWS 31 | MAY 21 | BIOSIS reloaded and enhanced with archival data |
| NEWS 32 | MAY 21 | TOXCENTER enhanced with BIOSIS reload |
| NEWS 33 | MAY 21 | CA/CAplus enhanced with additional kind codes for German patents |
| NEWS 34 | MAY 22 | CA/CAplus enhanced with IPC reclassification in Japanese patents |

NEWS EXPRESS NOVEMBER 10 CURRENT WINDOWS VERSION IS V8.01c, CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP), AND CURRENT DISCOVER FILE IS DATED 25 SEPTEMBER 2006.

Application No: 10/527,716

NEWS HOURS STN Operating Hours Plus Help Desk Availability
NEWS LOGIN Welcome Banner and News Items
NEWS IPC8 For general information regarding STN implementation of IPC 8

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FILE 'HOME' ENTERED AT 15:41:36 ON 05 JUN 2007

=> file reg

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|---|---|---|
| FULL ESTIMATED COST | 0.21 | 0.21 |

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STRUCTURE FILE UPDATES: 4 JUN 2007 HIGHEST RN 936539-19-4
DICTIONARY FILE UPDATES: 4 JUN 2007 HIGHEST RN 936539-19-4

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=>
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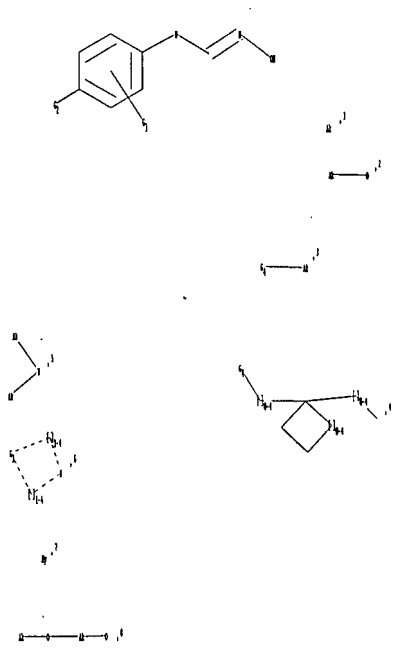

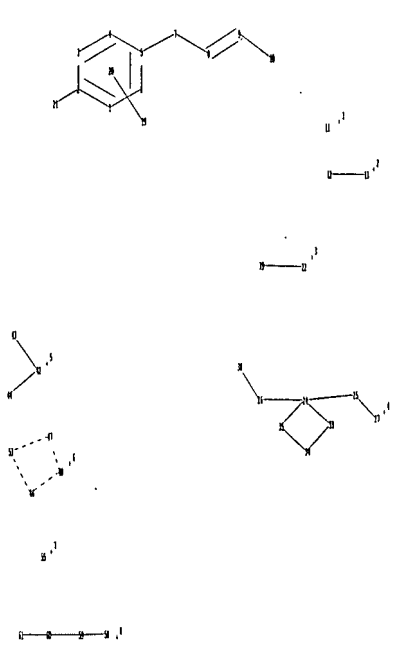

chain nodes :
7 8 9 10 11 12 13 19 21 22 25 26 27 38 39 42 43 44 55 58 59 60 61
ring nodes :
1 2 3 4 5 6 24 33 34 35 46 47 48 53
chain bonds :
2-21 5-7 7-8 8-9 9-10 12-13 22-39 24-25 24-26 25-27 26-38 42-43 42-44 58-59 59-60 60-61
ring bonds :
1-2 1-6 2-3 3-4 4-5 5-6 24-33 24-35 33-34 34-35 46-48 46-53 47-48 47-53

exact/norm bonds :
2-21 5-7 7-8 8-9 9-10 12-13 22-39 24-25 24-26 24-33 24-35 25-27 26-38
33-34 34-35 42-43 42-44 46-48 46-53 47-48 47-53 58-59 59-60 60-61
normalized bonds :
1-2 1-6 2-3 3-4 4-5 5-6

G1:H,X,[*1],[*2]

G2:[*3],[*4]

G3:C,O,S,N

G4:[*5],[*6],[*7],[*8]

Match level :
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:CLASS 8:CLASS 9:CLASS 10:CLASS
11:CLASS 12:CLASS 13:CLASS 19:CLASS 20:Atom 21:CLASS 22:CLASS 24:Atom
25:CLASS 26:CLASS 27:CLASS 33:Atom 34:Atom 35:Atom 38:CLASS 39:CLASS
42:CLASS 43:CLASS 44:CLASS 46:Atom 47:Atom 48:Atom 53:Atom 55:Atom 58:CLASS
59:CLASS 60:CLASS 61:CLASS
Element Count :
Node 11: Limited
C,C1-6

Node 12: Limited
C,C1-6

Node 22: Limited
C,C1-11

Node 55: Limited
O,O2
C,C4

Node 59: Limited
C,C1-4

Node 61: Limited
C,C1-4

L1 STRUCTURE UPLOADED

=> d
L1 HAS NO ANSWERS
L1 STR
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

Structure attributes must be viewed using STN Express query preparation.

=> s l1
SAMPLE SEARCH INITIATED 15:42:19 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 730 TO ITERATE

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100.0% PROCESSED      730 ITERATIONS                                1 ANSWERS
SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS:  ONLINE  **COMPLETE**
                        BATCH   **COMPLETE**
PROJECTED ITERATIONS:        12980 TO    16220
PROJECTED ANSWERS:               1 TO       80

L2            1 SEA SSS SAM L1

=> s l1 full
FULL SEARCH INITIATED 15:42:24 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED -    14688 TO ITERATE

100.0% PROCESSED    14688 ITERATIONS                               10 ANSWERS
SEARCH TIME: 00.00.01

L3           10 SEA SSS FUL L1

=> file zcaplus
COST IN U.S. DOLLARS                                SINCE FILE      TOTAL
                                                       ENTRY    SESSION
FULL ESTIMATED COST                                   172.10     172.31

FILE 'ZCAPLUS' ENTERED AT 15:42:30 ON 05 JUN 2007
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FILE COVERS 1907 - 5 Jun 2007  VOL 146 ISS 24
FILE LAST UPDATED: 4 Jun 2007  (20070604/ED)

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  This file contains CAS Registry Numbers for easy and accurate
  substance identification.

=> s l3
L4            4 L3

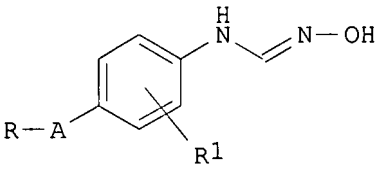
=> d ibib abs hitstr 1-4

L4   ANSWER 1 OF 4  ZCAPLUS  COPYRIGHT 2007 ACS on STN
ACCESSION NUMBER:         2005:1073980  ZCAPLUS
DOCUMENT NUMBER:          143:339637
TITLE:                    Agents for inhibition of 20-HETE-producing enzyme
                          containing N-hydroxyformamidine derivatives
INVENTOR(S):              Sato, Masakazu; Kakinuma, Hiroyuki; Amada, Hideaki
PATENT ASSIGNEE(S):       Taisho Pharmaceutical Co., Ltd., Japan
```

SOURCE: Jpn. Kokai Tokkyo Koho, 17 pp.
CODEN: JKXXAF
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|---|---|---|---|
| JP 2005272446 | A | 20051006 | JP 2005-44930 | 20050222 |
| PRIORITY APPLN. INFO.: | | | JP 2004-52724 A | 20040227 |

OTHER SOURCE(S): MARPAT 143:339637
GI

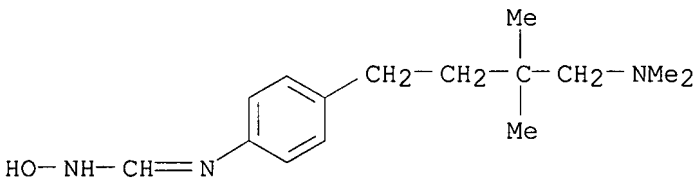


I

AB The invention relates to a 20-HETE (20-hydroxyeicosatetraenoic acid)-producing enzyme inhibitor characterized by containing N-hydroxyformamidine derivative I (R1 - H, C1-4 alkyl, C1-4 alkoxy, halogen, A = C1-10 alkylene, etc; R = N,N-diC1-6alkylamino, dioxanyl, C1-4 alkyl-substituted dioxanyl, C1-4 alkoxy C1-4 alkoxy, etc.) or its pharmaceutically acceptable salt as an active component. The agent is suitable for use for treatment of renal disease, cerebrovascular disease, and/or cardiovascular disease. For example, a compound N-[4-(4-N,N-dimethylamino-3,3-dimethyl-butyl)phenyl]-N'-hydroxyformamidine was prepared, and tested for its effect on human renal microsome-derived 20-HETE-producing enzyme in vitro.

IT 675583-50-3P 675583-51-4P 675583-52-5P
675583-53-6P 675583-54-7P 675583-55-8P
675583-56-9P 675583-57-0P
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(agents for inhibition of 20-HETE-producing enzyme containing N-hydroxyformamidine derivs.)

RN 675583-50-3 ZCAPLUS
CN Methanimidamide, N-[4-[4-(dimethylamino)-3,3-dimethylbutyl]phenyl]-N'-hydroxy- (9CI) (CA INDEX NAME)



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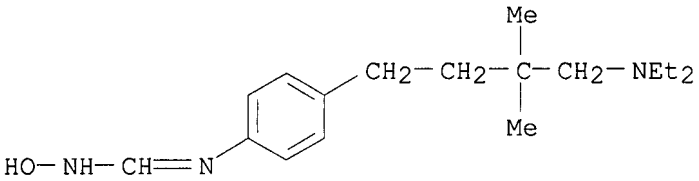

RN 675583-52-5 ZCAPLUS
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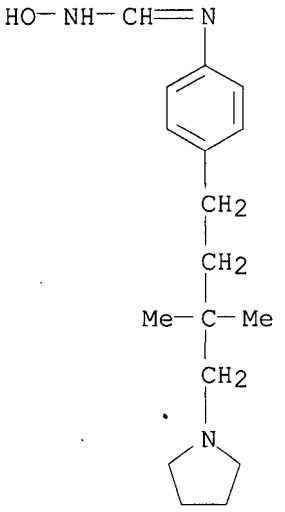

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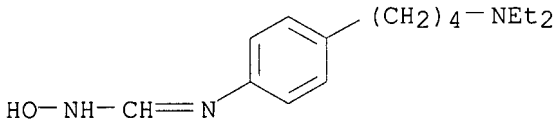
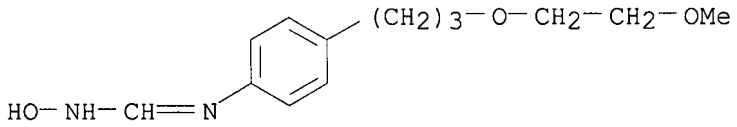


RN 675583-54-7 ZCAPLUS
CN Methanimidamide, N-hydroxy-N'-[4-[3-(2-methoxyethoxy)propyl]phenyl]- (9CI) (CA INDEX NAME)



RN 675583-55-8 ZCAPLUS
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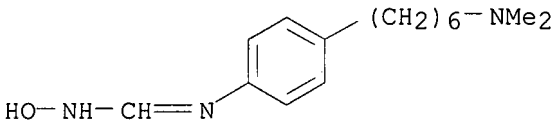

RN 675583-56-9 ZCAPLUS
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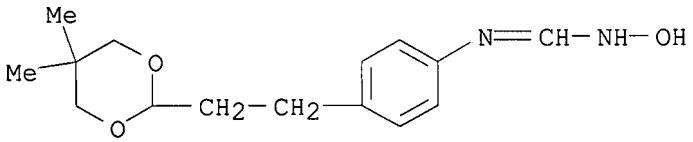

RN 675583-57-0 ZCAPLUS
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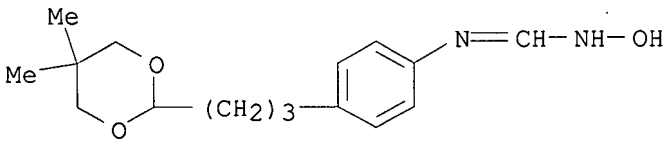

L4 ANSWER 2 OF 4 ZCAPLUS COPYRIGHT 2007 ACS on STN
ACCESSION NUMBER: 2004:252475 ZCAPLUS
DOCUMENT NUMBER: 140:287175
TITLE: Synthesis of N-hydroxyformamidine derivatives for 20-HETE inhibitor
INVENTOR(S): Sato, Masakazu; Kakinuma, Hiroyuki; Amada, Hideaki
PATENT ASSIGNEE(S): Taisho Pharmaceutical Co.,ltd., Japan
SOURCE: PCT Int. Appl., 25 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|---|---|---|---|
| WO 2004024677 | A1 | 20040325 | WO 2003-JP11603 | 20030911 |
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| RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| AU 2003262069 | A1 | 20040430 | AU 2003-262069 | 20030911 |

| | | | | | |
|---|---|---|---|---|---|
| CN 1681775 | A | 20051012 | CN 2003-821795 | | 20030911 |
| US 2006004078 | A1 | 20060105 | US 2005-527716 | | 20050311 |
| PRIORITY APPLN. INFO.: | | | JP 2002-266765 | A | 20020912 |
| | | | WO 2003-JP11603 | W | 20030911 |

OTHER SOURCE(S): MARPAT 140:287175


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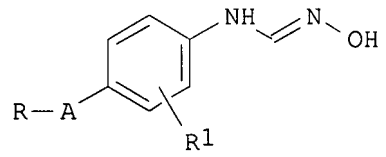

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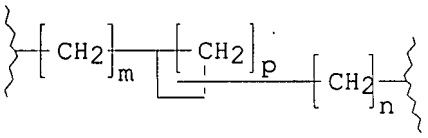

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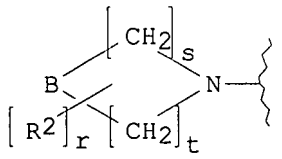
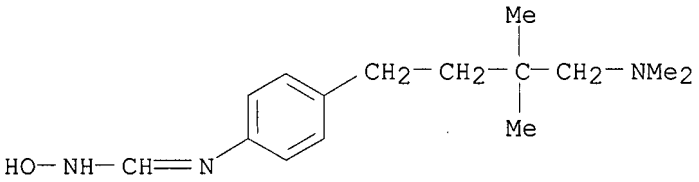


III


AB The patent relates to the synthesis of an N-hydroxyformamidine derivative I wherein R1 = H, C1-4 alkyl, C1-4 alkoxy, or halogeno; A = C1-10 alkylene or a group represented by the formula II (wherein m, n, and p = an integer of 0 to 4); and R = N,N-di(C1-6 alkyl)amino, dioxanyl, dioxanyl substituted by C1-4 alkyl, C1-4 alkoxy substituted by C1-4 alkoxy, or a group represented by the formula III (wherein s, t = 1 to 4; B = methylene, oxygen, sulfur, nitrogen, nitrogen substituted by C1-4 alkyl, nitrogen substituted by Ph, or nitrogen substituted by benzyl; R2 = H or C1-4 alkyl; and r is an integer of 0 to 2); and a pharmaceutically acceptable salt of the derivative Also provided is a medicine having an inhibitory activity against an enzyme producing 20-HETE, which relates to microvascular constrictive or dilative activity, cell proliferative activity, etc. in major organs such as the kidneys and cerebral blood vessels. Thus, N-[4-(4-N,N-dimethylamino-3,3-dimehtylbutyl)phenyl]-N'-hydroxyformamidine prepared from: reaction of dimethylammonium chloride, paraformaldehyde, isobutylaldehyde to form 3-dimethylamino-2,2-dimethylpropionaldehyde; reaction with di-Et benzylphosphonate to form (2,2-dimethyl-4-phenyl-3-butenyl)dimethylamine; hydrogenation to form (2,2-dimethyl-4-phenyl-3-butyl)dimethylamine; nitration to form a mixture of 2,2-dimethyl-4-(4-nitrophenyl-3-butyl)dimethylamine and 2,2-dimethyl-4-(2-nitrophenyl-3-butyl)dimethylamine; hydrogenation to form 4-(4-dimethylamino-3,3-dimethylbutyl)aniline; and reaction with N,N-dimethylforamide dimethylacetal was tested as inhibitor for 20-HETE and gave IG50 of 10.5 nM.

IT 675583-50-3P 675583-51-4P 675583-52-5P
675583-53-6P 675583-54-7P 675583-55-8P
675583-56-9P 675583-57-0P
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation of N-hydroxyformamidine derivs. for 20-HETE inhibitor)

RN 675583-50-3 ZCAPLUS
CN Methanimidamide, N-[4-[4-(dimethylamino)-3,3-dimethylbutyl]phenyl]-N'-hydroxy- (9CI) (CA INDEX NAME)



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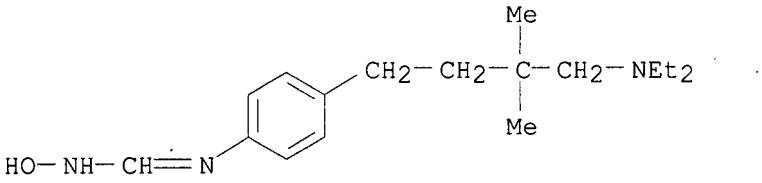

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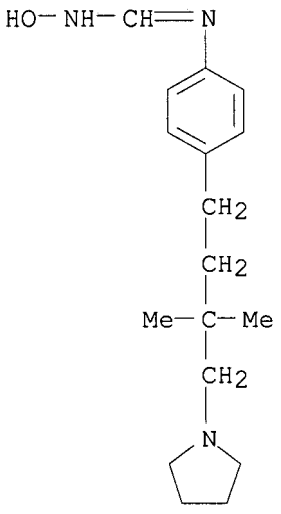

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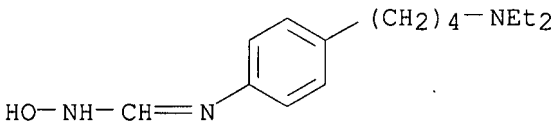
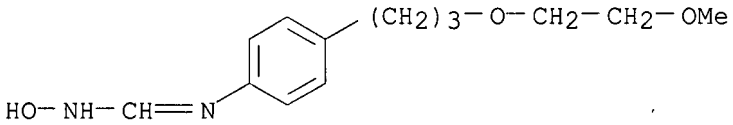
CN Methanimidamide, N-[4-[4-(diethylamino)butyl]phenyl]-N'-hydroxy- (9CI)

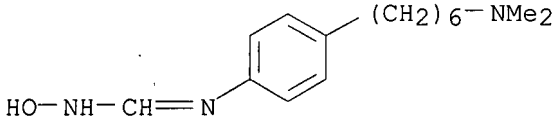
(CA INDEX NAME)



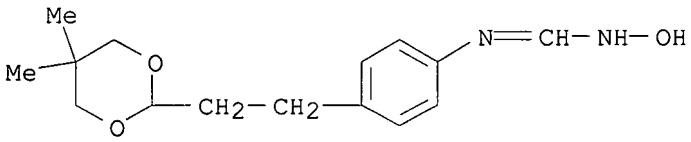
RN 675583-54-7 ZCAPLUS
CN Methanimidamide, N-hydroxy-N'-[4-[3-(2-methoxyethoxy)propyl]phenyl]- (9CI) (CA INDEX NAME)



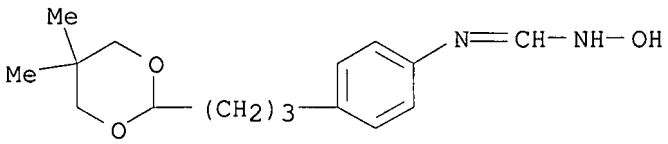
RN 675583-55-8 ZCAPLUS
CN Methanimidamide, N-[4-[6-(dimethylamino)hexyl]phenyl]-N'-hydroxy- (9CI) (CA INDEX NAME)



RN 675583-56-9 ZCAPLUS
CN Methanimidamide, N-[4-[2-(5,5-dimethyl-1,3-dioxan-2-yl)ethyl]phenyl]-N'-hydroxy- (9CI) (CA INDEX NAME)



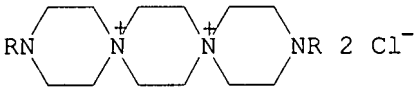
RN 675583-57-0 ZCAPLUS
CN Methanimidamide, N-[4-[3-(5,5-dimethyl-1,3-dioxan-2-yl)propyl]phenyl]-N'-hydroxy- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 3 OF 4 ZCAPLUS COPYRIGHT 2007 ACS on STN
ACCESSION NUMBER: 1981:174593 ZCAPLUS

DOCUMENT NUMBER: 94:174593
TITLE: Acetylformamidoxime derivatives and their antiblastic properties
AUTHOR(S): Poplevskaya, I. A.; Kurmangalieva, R. G.; Khalilova, S. F.; Abdullin, K. A.; Kudrina, I. K.; Kabiev, O. K.
CORPORATE SOURCE: USSR
SOURCE: Trudy Instituta Khimicheskikh Nauk, Akademiya Nauk Kazakhskoi SSR (1980), 52, 114-27.
CODEN: TIKNAG; ISSN: 0568-5087
DOCUMENT TYPE: Journal
LANGUAGE: Russian
GI

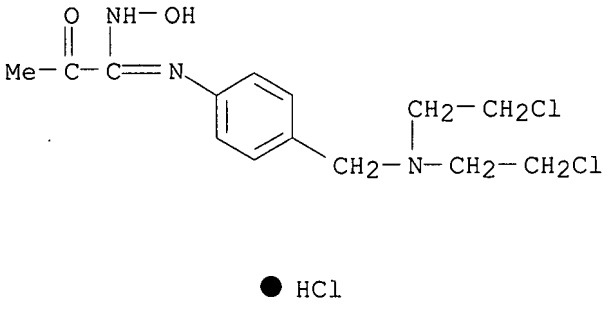


II

AB RN:CMeCCl:NOH [R = PhNH, HO (I)] condensed with HN(CH2CH2Cl)2 to give RN:CMeC(:NOH)N(CH2CH2Cl)2 (same R), isolated as the hydrochlorides. Analogous reaction of (HON:CClCMe:N)2 with p-H2NC6H4X [X = N(CH2CH2Cl)2, OMe] give [p-XC6H4NHC(:NOH)CMe:N]2 (same X), and I reacted with spiro diquaternary salt II (R = H) to give II [R = HON:CMeC(:NOH)]. These products and 13 related compds. were tested for antineoplastic activities; MeCOC(:NOH)NHC6H4N(CH2CH2Cl)2-p was the most active over the widest range of tumors.

IT 18237-82-6
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(antineoplastic activity of)

RN 18237-82-6 ZCAPLUS
CN Propanimidamide, N-[4-[[bis(2-chloroethyl)amino]methyl]phenyl]-N'-hydroxy-2-oxo-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

L4 ANSWER 4 OF 4 ZCAPLUS COPYRIGHT 2007 ACS on STN
ACCESSION NUMBER: 1968:402606 ZCAPLUS
DOCUMENT NUMBER: 69:2606
TITLE: Synthesis of α-amino-α-isonitrosoacetone derivatives
AUTHOR(S): Azerbaev, I. N.; Poplavskaya, I. A.; Kurmangalieva, R.

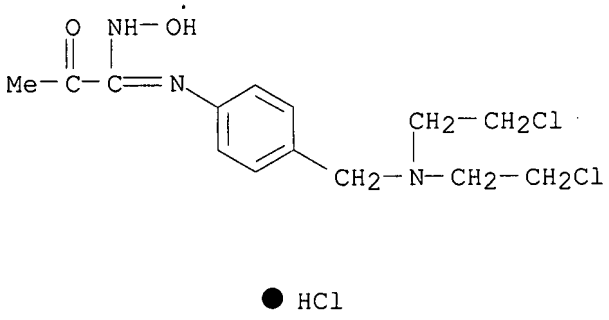
G.
CORPORATE SOURCE: Inst. Khim. Nauk, Alma-Ata, USSR
SOURCE: Zhurnal Organicheskoi Khimii (1968), 4(4), 590-4
CODEN: ZORKAE; ISSN: 0514-7492
DOCUMENT TYPE: Journal
LANGUAGE: Russian

AB Reaction of AcCCl:NOH (I) with amines of general formula RNH2, where R is p-(ClCH2CH2)2NC6H4CH2, p-(ClCH2CH2)2NC6H4, or p-(ClCH2CH2)2NCH2C6H4, in the presence of NEt3 as HCl acceptor gave AcC(NHR):NOH (II). p-(ClCH2CH2)2NCH2C6H4NH2 (III) was prepared by reduction over Pd/C of p-(ClCH2CH2)2NCH2C6H4NO2. An alternative route, LiAlH4 reduction of p-O2NC6H4CON(CH2CH2Cl)2, gave only a small amount of [p-(ClCH2CH2)2NCH2C6H4]2N2, the reduction of which to III was not attempted. Other II were prepared included (R in II given): 2-HOC6H4, 4-ClC6H4, 3,4-Cl2C6H3, 3-ClC6H4, 3,4-ClMeC6H3, and 2,5-MeClC6H3. However, reacting I with 2,4-Cl2C6H3NH2, 2,5-Cl2C6H3NH2, or 4-HO2CC6H4NH2 gave only the corresponding acetanilides.

IT 18237-82-6P
RL: SPN (Synthetic preparation); PREP (Preparation)
(preparation of)

RN 18237-82-6 ZCAPLUS

CN Propanimidamide, N-[4-[[bis(2-chloroethyl)amino]methyl]phenyl]-N'-hydroxy-2-oxo-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

=>

---Logging off of STN---

=>
Executing the logoff script...

=> LOG Y

| COST IN U.S. DOLLARS | SINCE FILE ENTRY | TOTAL SESSION |
|---|---|---|
| FULL ESTIMATED COST | 21.14 | 193.45 |
| DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) | SINCE FILE ENTRY | TOTAL SESSION |
| CA SUBSCRIBER PRICE | -3.12 | -3.12 |

Application No: 10/527,716

STN INTERNATIONAL LOGOFF AT 15:43:03 ON 05 JUN 2007